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Cooking fuel and respiratory symptoms among people living with HIV in rural Uganda

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ABSTRACT Household air pollution (HAP) and chronic HIV infection are each associated with significant respiratory morbidity. Little is known about relationships between HAP and respiratory symptoms among people living with HIV. The objective of this study was to investigate the relationship between cooking fuel type and chronic respiratory symptoms in study participants from the Uganda AIDS Rural Treatment Outcomes Study.

Study participants were enrolled at the time of antiretroviral therapy initiation and seen quarterly from 2005 to 2014 for health-focused questionnaires, CD4 count and HIV viral load. We used multivariable logistic regression and generalised estimating equations, with each study visit as a unit of observation, to investigate relationships between cooking fuel type and chronic respiratory symptoms.

We observed an association between cooking with firewood (*versus* charcoal) and chronic cough among HIV-infected females in rural Uganda (adjusted OR 1.41, 95% CI 1.00–1.99; $p=0.047$). We did not observe an association between cooking fuel type and respiratory symptoms among males (adjusted OR 0.88, 95% CI 0.47–1.63; $p=0.658$).

Associations between cooking fuel and chronic cough in this HIV-infected cohort may be influenced by sex-based roles in meal preparation. This study raises important questions about relationships between household air pollution, HIV infection and respiratory morbidity.



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This study raises important questions about relationships between air pollution, HIV and respiratory morbidity <http://ow.ly/zjsJ30arkI0>

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Introduction

More than 4 million deaths were attributed to household air pollution (HAP) worldwide in 2012. HAP predominantly results from the burning of biomass fuel for cooking and heating. Biomass fuels (firewood, charcoal, animal dung and crop residues) are highly polluting fuels used by more than half the world's population. Exposure to biomass fuels increases risk of chronic respiratory symptoms, particularly among females in low-income settings [1].

Chronic HIV infection is also independently associated with increased risk of chronic respiratory symptoms [2–6]. HIV and HAP are each hypothesised to cause pulmonary morbidity through systemic inflammatory pathways [7, 8], yet little is known about relationships between HAP and pulmonary morbidity among people living with HIV. It is plausible, for example, that HAP causes disproportionate pulmonary morbidity among people living with HIV due to similar effects on systemic inflammation. We analysed data from the Uganda AIDS Rural Treatment Outcomes Study (UARTO), a longitudinal cohort of HIV-infected persons taking antiretroviral therapy (ART), to estimate associations between HAP and respiratory symptoms among an HIV-infected population in Uganda.

Materials and methods

Adults aged >18 years were enrolled at the time of ART initiation during 2005–2014. Participants were seen quarterly to complete questionnaires and undergo phlebotomy for CD4 count and HIV-1 RNA viral load. At each visit, participants were asked about a history of cough or dyspnoea in the past 30 days. Those who reported cough were asked if the duration was >4 weeks. Participants who confirmed respiratory symptoms in the past 30 days were asked how bothersome each symptom was, on a four-point scale (“not at all”, “a little”, “moderately” and “a lot”). In addition, participants completed an annual questionnaire, in which they were asked “What is the main fuel used for cooking?”

We fit multivariable logistic regression models using generalised estimating equations and an exchangeable matrix structure with each study visit as a unit of observation to measure associations between respiratory symptoms and cooking fuel. Our outcomes of interest were 1) any self-reported cough or dyspnoea and 2) a self-reported cough of ≥ 4 weeks' duration (“chronic cough”). Our exposure of interest was self-reported cooking fuel type: firewood or charcoal (which were reported in 97% of study visits). Models were adjusted for known correlates of respiratory symptoms, including age, smoking status (current/former/never), occupation, household asset index, CD4 count and viral load. We explored prespecified interactions between cooking fuel type and sex because Ugandan females do most of the cooking and could have higher exposure to biomass fuels, and used postestimation margins to compare the adjusted proportion of participants who reported each outcome by cooking fuel type. We tested the significance of global categorical model coefficients (e.g. CD4 count strata) using a Wald test.

Data were analysed using Stata 13 (StataCorp, College Station, TX, USA). All study procedures were approved by Partners HealthCare (Boston, MA, USA) and the Mbarara University of Science and Technology (Mbarara, Uganda), and all participants gave written informed consent.

Results

Out of 762 participants enrolled in UARTO, 742 (97%) completed at least one questionnaire on respiratory symptoms and cooking fuel use. Eight (1%) had missing data for at least one covariate at each study visit and were excluded. Overall, 734 participants contributed 2757 study visits (median 20 visits) and were followed for a median (interquartile range (IQR)) 5 (2–7) years (table 1). At the time of study enrolment, 70% (n=511) were female, median (IQR) age was 34 (28–40) years, median (IQR) CD4 count was 167 (95–260) μL^{-1} , 23% (n=172) had ever smoked and 58% (n=428) reported firewood as the primary cooking fuel.

Respiratory symptoms were commonly reported during the observation period. Study participants reported cough or dyspnoea in the past 30 days or chronic cough at 30% (n=827) and 10% (n=287) of visits, respectively (figure 1). Among those who self-reported respiratory symptoms, 53% (n=200) of those reporting cough and 48% (n=92) of those reporting dyspnoea quantified the symptom burden as “a little”.

In multivariable models adjusted for age, occupation, household asset ownership index (a proxy for socioeconomic status), smoking status, cooking fuel type, CD4 count and viral load, cooking with firewood as compared to charcoal was associated with increased odds of chronic cough among females (adjusted OR 1.41, 95% CI 1.00–1.99; $p=0.047$). This corresponds to a 3.0% excess (95% CI 0.1–5.9%) in the proportion of visits when chronic cough was reported for those using firewood (12%) *versus* charcoal (9%). We found no relationships between respiratory symptoms and cooking fuel type among males (adjusted OR 0.88, 95% CI 0.47–1.63; $p=0.658$). In multivariable models adjusted for age, sex, occupation, smoking status and cooking fuel type, chronic cough was additionally associated with lower socioeconomic

TABLE 1 Summary characteristics for participants at the enrolment study visit

	Total cohort	Males	Females	p-value [#]
Subjects n	734	223	511	
Age years	34 [28–40]	37 [32–43]	32 [27–38]	<0.001
Smoking status[¶]				<0.001
Current	90 [12]	62 [28]	28 [5]	
Former	82 [11]	44 [20]	37 [7]	
Never	562 [77]	116 [52]	446 [87]	
Household asset ownership index[*]				NS
Lowest quartile	235 [32]	62 [28]	173 [34]	
Second quartile	173 [24]	57 [26]	116 [23]	
Third quartile	177 [24]	58 [26]	119 [23]	
Highest quartile	140 [19]	42 [19]	98 [19]	
High-risk occupation[§]	237 [32]	117 [52]	120 [23]	<0.001
Cooking fuel type				0.001
Charcoal	297 [40]	70 [31]	227 [44]	
Firewood	428 [58]	149 [67]	279 [55]	
Self-reported pulmonary disease history				
Pulmonary tuberculosis	60 [8]	26 [12]	34 [7]	0.02
Pneumocystis pneumonia	38 [5]	14 [6]	24 [5]	NS
Cryptococcal pneumonia	4 [1]	2 [1]	2 [<1]	NS
CD4 count				NS
<100 cells· μL^{-1}	198 [27]	73 [33]	125 [24]	
100–349 cells· μL^{-1}	441 [60]	124 [56]	317 [62]	
350–499 cells· μL^{-1}	59 [8]	18 [8]	41 [8]	
≥ 500 cells· μL^{-1}	34 [5]	7 [3]	27 [5]	
Viral load				0.01
$\leq 10\,000$ copies· mL^{-1}	79 [11]	14 [6]	65 [13]	
$> 10\,000$ copies· mL^{-1}	649 [88]	206 [92]	443 [87]	
Study visits per participant n	20 [7–28]	16 [7–28]	21 [8–28]	NS
Observation time years	5 [2–7]	4 [2–7]	5 [2–7]	NS

Data are presented as n [%] or median (interquartile range), unless otherwise stated. [#]: p-values for comparisons between sexes, calculated using t-test for normally distributed variables, Wilcoxon rank sum testing for non-normally distributed variables and Chi-squared testing for dichotomous and categorical variables; [¶]: never-smokers defined as those who did not smoke prior to cohort enrolment and continued not to smoke; former smokers defined as those who smoked prior to cohort enrolment, but did not smoke throughout the duration of the study; current smokers defined as those who smoked at cohort enrolment and continued to smoke throughout the duration of the study; ^{*}: constructed following the method of FILMER and PRITCHETT [9], using the following binary indicator variables: home ownership; land ownership; livestock ownership; presence of the following items (yes/no) in the household: iron, stove, fridge, phone, motorcycle, clock, bed, sofa, bike, television, lantern, cupboard, shoes, car, radio, mattress or electricity; characteristics of the home: flushing toilet, cement wall, cement floor or piped water; [§]: includes farmer, cook, motorcyclist, brick worker, builder, carpenter, cattle keeper, dry cleaner and sand miner. NS: nonsignificant.

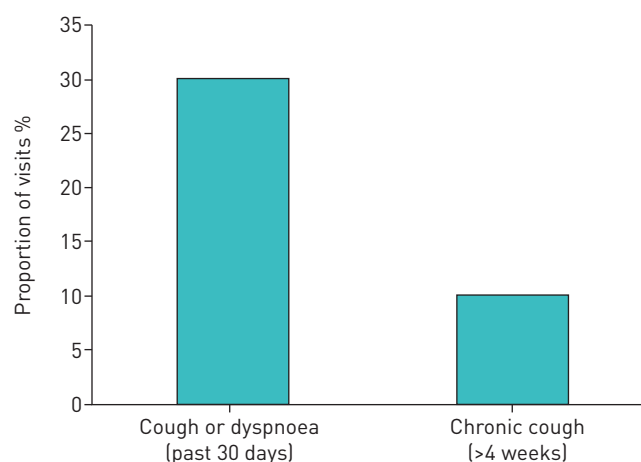


FIGURE 1 Frequency of self-reported respiratory symptoms at study visits as a proportion of total study visits.

status ($p=0.01$), lower CD4 count ($p<0.01$) and increased HIV viral load ($p<0.001$) (table 2 and figure 2). Interaction effects between cooking fuel type and degree of immunosuppression (CD4 count and HIV-1 viral load) were not statistically significant ($p>0.3$ for all interaction terms).

Discussion

We identified an association between use of firewood for cooking (as compared to charcoal) and chronic cough among HIV-infected females in rural Uganda, adjusted for indices of HIV disease stage (CD4 count or viral load). To our knowledge, this is the first investigation of the association between cooking fuel and respiratory symptoms in an HIV-infected cohort. The results of this study are consistent with published literature on the respiratory effects of HAP in the general population, which has been associated with chronic respiratory symptoms, impaired pulmonary function, respiratory infections and lung cancer in HIV-uninfected cohorts, particularly among females [10]. Furthermore, prior work has demonstrated that chronic cough poses a greater burden on health-related quality of life for females than males [11]. Although it is plausible that they are related, the consequences of HAP on respiratory symptoms should not necessarily be assumed to be similar between HIV-infected and uninfected populations. Either synergistic or attenuated effects of HAP on the pulmonary health of HIV-infected individuals can be postulated based on formative data in the field. For example, HAP and HIV each cause systemic inflammation [7, 8], which may result in an additive or multiplicative effect of HAP on respiratory pathology among HIV-infected individuals, compared to HIV-uninfected populations. Similarly, HIV-infected individuals are more susceptible to respiratory infections due to immune compromise, resulting in higher incidences of pulmonary tuberculosis, pneumocystis and bacterial pneumonias [12]. Thus, any incremental increases in respiratory symptom burden caused by HAP exposure among HIV-infected individuals must also take into account underlying lung disease and whether it may attenuate the absolute effects of HAP on lung health. By demonstrating a relationship between HAP and respiratory symptoms, our data provide a rationale for further quantifying these complex associations among HIV-infected populations.

TABLE 2 Adjusted odds ratios [95% CI] for predictors of cough of ≥ 4 weeks' duration

	Total cohort	Males	Females
Observations n	2751	778	1973
Age			
<30 years	Ref.	Ref.	Ref.
30–39 years	0.86 [0.61–1.21]	0.74 [0.32–1.76]	0.85 [0.59–1.24]
40–49 years	0.88 [0.59–1.33]	0.82 [0.32–2.09]	0.91 [0.58–1.43]
≥ 50 years	1.39 [0.80–2.41]	2.78 [1.01–7.62]	0.72 [0.32–1.62]
Female	1.18 [0.83–1.69]		
High-risk occupation[#]	0.78 [0.57–1.06]	0.72 [0.42–1.25]	0.77 [0.53–1.13]
Household asset ownership index			
Lowest quartile	Ref.	Ref.	Ref.
Second quartile	0.96 [0.68–1.35]	1.00 [0.50–2.01]	0.95 [0.64–1.41]
Third quartile	0.61 [0.42–0.88]	0.47 [0.21–1.04]	0.67 [0.44–1.02]
Highest quartile	0.62 [0.42–0.91]	0.55 [0.24–1.27]	0.64 [0.41–1.01]
Smoking status			
Never	Ref.	Ref.	Ref.
Former	0.95 [0.62–1.45]	0.95 [0.49–1.86]	0.96 [0.54–1.69]
Current	1.11 [0.65–1.89]	0.83 [0.39–1.79]	1.75 [0.81–3.78]
CD4 count			
<100 cells· μL^{-1}	2.16 [1.24–3.77]	3.20 [0.89–11.42]	1.75 [0.92–3.32]
100–349 cells· μL^{-1}	1.20 [0.76–1.90]	1.16 [0.36–3.73]	1.21 [0.74–2.00]
350–499 cells· μL^{-1}	1.01 [0.61–1.68]	1.40 [0.41–4.82]	0.90 [0.52–1.58]
≥ 500 cells· μL^{-1}	Ref.	Ref.	Ref.
Viral load			
Undetectable	Ref.	Ref.	Ref.
$\leq 10\,000$ copies· mL^{-1}	1.07 [0.69–1.67]	0.33 [0.08–1.40]	1.38 [0.85–2.22]
$> 10\,000$ copies· mL^{-1}	4.59 [3.30–6.38]	6.48 [3.26–12.89]	4.37 [2.97–6.42]
Cooking source			
Charcoal	Ref.	Ref.	Ref.
Firewood	1.25 [0.93–1.69]	0.90 [0.48–1.67]	1.41 [1.00–1.99] [¶]

[#]: includes farmer, cook, motorcyclist, brick worker, builder, carpenter, cattle keeper, dry cleaner and sand miner; [¶]: $p=0.047$; p -value for interaction between cooking source and sex $p=0.21$.

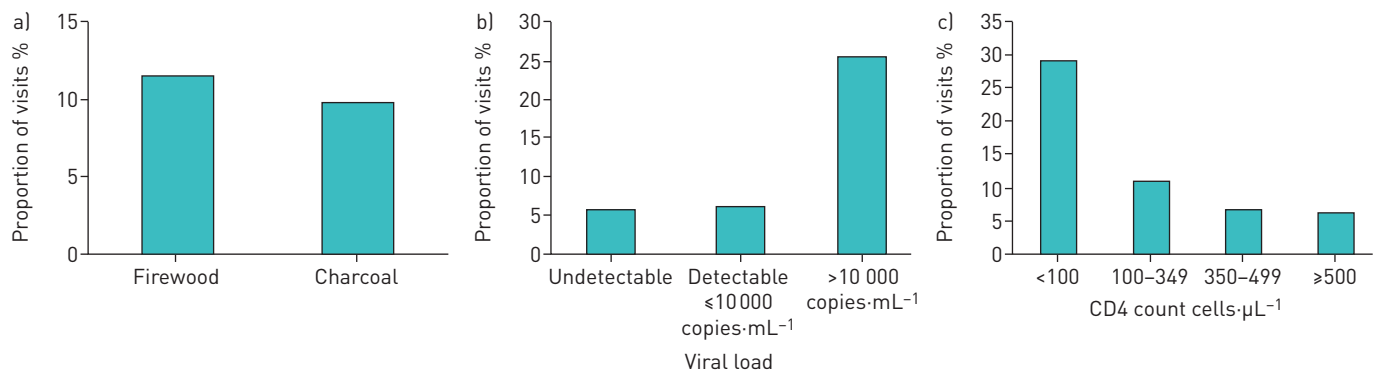


FIGURE 2 a) Proportion of study visits with reported chronic cough among females, stratified by cooking fuel type; b) proportion of study visits with reported chronic cough stratified by viral load; c) proportion of study visits with reported chronic cough stratified by CD4 count.

While there are no comparable studies among HIV-infected populations in sub-Saharan Africa, two recent studies measured the odds of cough among biomass-exposed populations in Malawi [13] and Nigeria [14]. Similar to our work, both studies identify increased odds of cough among females cooking with firewood. Urban and rural Malawian females who cooked with firewood as compared to charcoal had >300% increased odds of cough, and rural Nigerian females cooking with firewood as compared to kerosene gas had >400% increased odds of cough. Neither study reported data on HIV prevalence. Two notable differences might explain the increased odds of cough noted in those studies compared to our work. Firstly, the Malawian study population was 42% peri-urban and focused on females with primary cooking responsibilities, each of which may have led to increased pollutant exposure among study participants. Secondly, the Nigerian study compared firewood to a clean-burning fuel, whereas we compared firewood to the only other fuel prevalent regionally: charcoal, a biomass fuel.

The elevated risk in models restricted to females, but not males, may be due to the role of females in cooking in the region [10]. The increased chronic cough risk among those using firewood *versus* charcoal may be related to airborne endotoxin inhalation [15], which is associated with chronic respiratory symptoms and accelerated pulmonary function decline [16].

Chronic respiratory symptoms, including cough and dyspnoea, are prevalent self-reported conditions in established HIV cohorts in high-income settings, with comparable proportions to those reported here [2, 4]. Markers of inadequate HIV viral control were associated with chronic cough in our cohort, in keeping with published literature in several large US-based HIV-infected cohorts that have shown lower CD4 count and higher HIV-1 viral load to be independent risk factors for chronic respiratory symptoms [2, 4, 6]. In addition, respiratory symptoms in these cohorts are associated with impaired pulmonary function, postulated to be a result of prior pulmonary opportunistic infections and/or chronic inflammation-associated obstructive lung disease. Our study was limited to an HIV-infected population, and we are therefore unable to assess the relative contribution of HIV infection *per se* to the chronic cough burden. Importantly, HAP and HIV are each associated with chronic respiratory symptoms, so an essential unanswered question raised by our work is whether there are interactive effects between HIV infection and HAP that exacerbate chronic respiratory disease risk.

We also identified an association between chronic cough and lower socioeconomic status, as measured by the household asset index [9]. Our results are in accordance with published literature on associations with chronic cough, in which lower socioeconomic status portends increased risk of chronic productive cough [17, 18] and chronic bronchitis [19–21].

The main strength of this analysis is the reliance on a large, longitudinal study of HIV-infected persons in rural Uganda with rare missing data or losses from observation. The results of our work can be applied to HIV-infected people living in environments with high biomass exposure.

Our study has several limitations. Firstly, cooking fuel type and respiratory symptoms are self-reported and subject to misclassification. However, any misclassification is likely to be noninformative and bias towards the null. Secondly, we compare firewood to another biomass fuel source (charcoal), rather than a cleaner burning fuel. However, this probably underestimates any true association between biomass fuel exposure in general and chronic cough in this population. Furthermore, >80% of sub-Saharan Africa is exposed to biomass smoke [22]. Comparing the two regionally prevalent biomass sources more accurately reflects the local biomass exposure patterns, increases the generalisability of our results to the HIV-infected sub-Saharan African community and provides critical knowledge to inform risk-reduction practices while global access to

cleaner energy sources is evolving. Thirdly, no data were collected on the quantity of cooking fuel used or the participant's role in cooking activities, and self-reported cooking fuel type is a blunt instrument for quantification of household air pollution exposure. We partially accounted for this with models stratified by sex to account for presumed increased exposure among females, but acknowledge that future studies that comprehensively quantify biomass and other environmental exposures are necessary. Lastly, pulmonary function testing was not performed in this cohort. This limits inferences between chronic respiratory symptoms and pulmonary function.

Conclusions

In summary, we describe an increased risk of chronic cough associated with firewood use among HIV-infected females in rural Uganda. The epidemics of household air pollution and HIV converge in sub-Saharan Africa, where chronic respiratory disease is a leading cause of death and disability [23]. Much chronic respiratory disease is preventable. Our results signal a need to pursue further investigations to elucidate the public health implications of relationships between HIV, biomass exposure and chronic respiratory disease.

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